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Substitute for form 1449/PTO				Complete if Known	
INFORMATION DISCLOSURE STATEMENT BY APPLICANT <i>(Use as many sheets as necessary)</i>				Application Number	10/632,187
				Filing Date	July 30, 2003
				First Named Inventor	Jürgen Engel
				Art Unit	1628
				Examiner Name	Shirley Gembeh
Sheet	1	of	6	Attorney Docket Number	ZEN-015-NP (Old 103832-477-NP)

NON PATENT LITERATURE DOCUMENTS				
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		ASCO Abstract 2006 - Journal of Clinical Oncology, 2006 ASCO Annual Meeting Proceedings (Post-Meeting Edition)m Vol. 24, No. 18S (June 20 Supplement), 2006.		
		ASCO Abstract 2009 - Journal of Clinical Oncology, 2009 ASCO Annual Meeting Proceedings (Post-Meeting Edition)m Vol. 27, No. 15S (May 20 Supplement), 2009:4081.		
		AYALA, GUSTAVO, et al., Bortezomib-Mediated Inhibition of Steroid Receptor Coactivator-3 Degradation Leads to Activated Akt, Clin Cancer Res, November 15, 2008, 7511-7518, 14(22).		
		CATLEY LAURENCE, et al., Alkyl Phospholipid Perifosine Induces Myeloid Hyperplasia in a Murine Myeloma Model, Experimental Hematology, 35 (2007), 1038-1046.		
		CHIARINI, F., et al., The novel Akt Inhibitor, Perifosine, Induces Caspase-Dependent Apoptosis and Downregulates P-glycoprotein Expression in Multidrug-Resistant Human T-acute Leukemia Cells by a JNK-dependent Mechanism, Leukemia (2008) 22, 1106-1116.		
		CIRSTEA, DIANA, et al., Dual Inhibition of Akt/Mammalian Target of Rapamycin Pathway by Nanoparticle Albumin-Bound-Rapamycin and Perifosine Induces Antitumor Activity in Multiple Myeloma, Mol Cancer Ther, April 2010, 963-975, 9(4).		
		CRUL, M., et al. Phase I and Pharmacological Study of Daily Oral Administration of Perifosine (D-21266) in Patients with Advanced Solid Tumours, European Journal of Cancer, 38 (2002), 1615-1621.		
		DASMAHAPATRA GIRIJA P., et al., In vitro Combination Treatment with Perfosine and UCN-01 Demonstrates Synergism Against Prostate (PC-3) and Lung (A549) Epithelial Adenocarcinoma Cell Lines, Clin Cancer Res, August 1, 2004, 5242-5252, Vol. 10.		
		DAVID, E., et al., Perifosine Synergistically EnhancesTRAIL-Induced Myeloma Cell Apoptosis via Up-Regulation of Death Receptors, Clin Cancer Res 2008:14(16) August 15, 2008, 5090-5098.		
		DOGAN, S. SERDAR, Ocular Side Effects Associated with Imatinib Mesylate and Perifosine for Gastrointestinal Stromal Tumor, Hematol Oncol. Clin N Am, 23 (2009) 109-114.		

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		ELLIS, MATTHEW J., et al., "PIKing" the Winner for Phosphatidylinositol 3-Kinase Inhibitors in ErbB2-Positive Breast Cancer: Let's Not "PTENed" It's Easy!, Clin Cancer Res, October 1, 2007, 5661-5662, 13(19).	
		ENGEL, JÖRG B., et al., Induction of Programmed Cell Death By Inhibition of AKT with the Alkylphosphocholine Perifosine in In Vitro Models of Platinum Sensitive and Resistant Ovarian Cancers, Arch Gynecol Obstet, doi:10.1007/s00404-010-1457-6; published online April 20, 2010.	
		ENGEL, JÖRG B., et al., Perifosine Inhibits Growth of Human Experimental Endometrial Cancers By Blockade of AKT Phosphorylation, Eur J. Obstet. Gynecol (2008), doi:10.1016/j.ejogrb.2008.06.007.	
		ERNST, D. SCOTT, et al. Phase II Study of Perifosine in Previously Untreated Patients with Metastatic Melanoma, Investiational New Drugs, 23:569-576, 2005.	
		FESTUCCIA, CLAUDIO, et al., Akt Down-Modulation Induces Apoptosis of Human Prostate Cancer Cells and Synergizes with EGFR Tyrosine Kinase Inhibitors, The Prostate, 58:965-974 (2008).	
		FLORYK, DANIEL, et al., Perifosine Induces Differentiation and Cell Death in Prostate Cancer Cells, Cancer Letters, 266 (2008) 216-226.	
		FOMCHENKO, ELENA I., et al., Mouse Models of Brain Tumors and Their Applications in Preclinical Trials, Clin Cancer Res., 2006;12(18) September 15, 2006; 5288-5297.	
		FU, LEI, et al., Perifosine Inhibits Mammalian Target of Rapamycin Signaling through Facilitating Degradation of Major Components in the mTOR Axis and Induces Autophagy, Cancer Res, December 1, 2009, 8967-8976, 69(23).	
		GAJATE, CONSUELO, et al., Edelfosine and Perifosine Induce Selective Apoptosis in Multiple Myeloma by Recruitment of Death Receptors and Downstream Signaling Molecules into Lipids Rafts, Blood, January 15, 2007, 711-719, Vol. 109, Number 2.	
		GILLS, JOELL, et al., Perifosine: Update on a Novel Akt Inhibitor, Current Oncology Reports-Evolving Therapies, 11:102-110.	

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		HARVEY, R. DONALD, et al., P13 Kinase/AKT Pathway as a Therapeutic Target in Multiple Myeloma, Future Oncol., (2007) 3(6), 639-647.		
		HIDESHIMA, TERU, et al., Perifosine, An Oral Bioactive Novel Alkylphospholipid, Inhibits Akt and Induces In Vitro and In Vivo Cytotoxicity in Human Multiple Myeloma Cells, Blood First Edition Paper, January 17, 2006, 1-34, doi 10.1182/blood-2005-08-3434.		
		HIDESHIMA, TERU, et al., Inhibition of Akt Induces Significant Downregulation of Survivin and Cytotoxicity in Human Multiple Myeloma Cells, British Journal of Haematology, 2007, 138, 783-791.		
		HUSTON, ALISSA, et al., Targeting and Heat Shock Protein 90 Produces Synergistic Multiple Myeloma Cell Cytotoxicity in the Bone Marrow Microenvironment, Clin Cancer Res, February 1, 2008, 865-874, 14(3).		
		JENDROSSEK, V., et al., Membrane Targeted Anticancer Drugs: Potent Inducers of Apoptosis and Putative Radiosensitisers, Curr. Med. Chem. - Anti-Cancer Agents, 2003, 3, 343-353.		
		KNOWLING, M., et al., A Phase II Study of Perifosine (D-21226) in Patients with Previously Untreated Metastatic or Locally Advanced Soft Tissue Sarcoma: A National Cancer Institute of Canada Clinical Trials Group Trial, Invest New Drugs, (2006), 24:435-439.		
		KODACH, LIUDMILA L., vilacein Synergistically Increases 5-fluorouracil Cytotoxicity, Induces Apoptosis and Inhibits Akt-mediated Signal Transduction in Human Colorectal Cancer Cells, Carcinogenesis, 2006, Vol. 27, No. 3, 508-516.		
		KONDAPAKA, SUDHIR B., Perifosine, A Novel Alkylphospholipid, Inhibits Protein Kinase B Activation, Molecular Cancer Therapeutics, 1093-1103.		
		KONSTANTINOV, SPIRO M., et al., BCR-ABL Influences the Antileukaemic Efficacy of Alkylphosphocholines, British Journal of Haematology, 1999, 107, 365-374.		
		KONSTANTINOV, SPIRO M., et al., Human Urinary Bladder Carcinoma Cell Lines Respond to Treatment with Alkylphosphocholines, Cancer Letters, 1999, 144, 153-160.		

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		KONSTANTINOV, SPIRO M., et al., Alkylphosphocholines: Effects on Human Leukemic Cell Lines and Normal Bone Marrow Cells, Int. J. Cancer, 1998, 77, 778-786.	
		KUMAR, ANIL, et al., The Alkylphospholipid Perifosine Induces Apoptosis and p21-Mediated Cell Cycle Arrest in Medulloblastoma, Mol Cancer Res, November 2009, 1813-1821, 7(11).	
		LEIGH, NATASHA B., et al., A Phase 2 Study of Perifosine in Advanced or Metastatic Breast Cancer, Breast Cancer Res Treat (2008), 108:87-92.	
		LELEU, XAVIER, et al., Targeting NF- κ B in Waldenstrom Macroglobulinemia, Blood, May 15, 2008, 111(10), 5068-5077.	
		LI, X., et al., Enhancement of Antitumor Activity of the Anti-EGF Receptor Monoclonal Antibody Cetuximab/C225 by Perifosine in PTEN-deficient Cancer Cells, Oncogen, (2005), 1-11.	
		LOPICCOLO, JACLYN, et al., Targeting the P13K/Akt/mTOR Pathway: Effective Combinations and Clinical Considerations, Drug Resistance Updates, 11 (2008) 32-50.	
		MITSIADES, CONSTANTIN S., et al., Emerging Treatments for Multiple Myeloma: Beyond Immunomodulatory Drugs and Bortezomib, Seminars in Hematology, April 2009, Vol. 46, No. 2, 166-175.	
		MOMOTA, HIROYUKI, et al., Perifosine Inhibits Multiple Signaling Pathways in Glial Progenitors and Cooperates With Temozolomide to Arrest Cell Proliferation in Gliomas In Vivo, Cancer Res, August 15, 2005, 65(16), 7429-7435.	
		NELSON, EC, et al., Inhibition of Akt Pathways in the Treatment of Prostate Cancer, Prostate Cancer and Prostatic Diseases, (2007), 10, 331-339.	
		NYÅKERN, MARIA, et al., Synergistic Induction of Apoptosis in Human Leukemia T Cells by the Akt Inhibitor Perifosine and Etoposide Through Activation of Intrinsic and Fas-mediated Extrinsic Cell Death Pathways, Mol Cancer Ther, June 2006, 5(6), 1559-1570.	

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		PAPA, V., et al., Proapoptotic Activity and Chemosensitizing Effect of the Novel Akt Inhibitor Perifosine in Acute Myelogenous Leukemia Cells, Leukemia, (2008), 22, 147-160.	
		PATEL, VYOMESH, Perifosine, A Novel Alkylphospholipid, Induces p21WAF1 Expression in Squamous Carcinoma Cells Through a P53-independent Pathway, Leading to Loss in Cyclin-dependent Kinase Activity and Cell Cycle Arrest, Cancer Research, 62, 1401-1409, 2002.	
		PORTA, CAMILLO, et al., Phosphatidylinositol-3-Kinase/Akt Signaling Pathway and Kidney Cancer, and the therapeutic Potential of Phosphatidylinositol-3-Kinase/Akt Inhibitors, The Journal of Urology, December 2009, Vol. 182, 2569-2577.	
		RAHMANI, MOHAMED, et al., Coadministration of Histone Deacetylase Inhibitors and Perifosine Synergistically Induces Apoptosis in Human Leukemia Cells Through Akt and ERK1/2 Inactivation and the Generation of Ceramide and Reactive Oxygen Species, Cancer Res, 2005, 2422-2432.	
		TAZAARI, PIER LUIGI, et al., Synergistic Proapoptotic Activity of Recombinant TRAIL Plus the Akt Inhibitor Perifosone in Acute Myelogenous Leukemia Cells, Cancer Res, November 15, 2008, 68 (22), 9394-9403.	
		UMMERSEN et al., A Phase I Trial of Perifosine (NSC 639966) on a Loading Dose/Maintenance Dose Schedule in Patients with Advanced Cancer, Clinical Cancer Research, Vol. 10, 7450-7456, November 15, 2004.	
		UNGER, CLEMENS, et al., First-Time-In-Man and Pharmacokinetic Study of Weekly Oral Perifosine in Patients with Solid Tumours, European Journal of Cancer, 46 (2010), 920-925.	
		VINALL, RUTH L., et al., Combination Treatment of Prostate Cancer Cell Lines with Bioactive Soy Isoflavones and Perifosine Causes Increased Growth Arrest and/or Apoptosis, Clin Cancer Res, October 15, 2007, 13(20), 6204-6216.	
		VINK, STEFAN R., et al., Tumor and Normal Tissue Pharmacokinetics of Perifosine, An Oral Anti-Cancer Alkylphospholipid, Investigational New Drugs, 23, 2005, 279-286.	
		VINK, STEFAN R., et al., Phase I and Pharmacokinetic Study of Combined Treatment With Perifosine and Radiation in Patients With Advanced Solid Tumours, Radiotherapy and Oncology, 80, 2006, 207-213.	

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